

## STN Columbus

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NEWS 7 DEC 21 IPC search and display fields enhanced in CA/Caplus with the  
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NEWS 8 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/  
USPAT2  
NEWS 9 JAN 13 IPC 8 searching in IFIPAT, IFIUDB, and IFICDB  
NEWS 10 JAN 13 New IPC 8 SEARCH, DISPLAY, and SELECT enhancements added to  
INPADOC  
NEWS 11 JAN 17 Pre-1988 INPI data added to MARPAT  
NEWS 12 JAN 17 IPC 8 in the WPI family of databases including WPIFV  
NEWS 13 JAN 30 Saved answer limit increased  
NEWS 14 JAN 31 Monthly current-awareness alert (SDI) frequency  
added to TULSA

NEWS EXPRESS FEBRUARY 15 CURRENT VERSION FOR WINDOWS IS V8.01a,  
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.  
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<http://download.cas.org/express/v8.0-Discover/>

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NEWS WWW CAS World Wide Web Site (general information)

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FILE 'HOME' ENTERED AT 11:32:18 ON 18 FEB 2006

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|----------------------|---------------------|------------------|
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## STN Columbus

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DICTIONARY FILE UPDATES: 16 FEB 2006 HIGHEST RN 874442-49-6

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*
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* effective March 20, 2005. A new display format, IDERL, is now    *
* available and contains the CA role and document type information. *
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L1 STRUCTURE UPLOADED

=> s l1

SAMPLE SEARCH INITIATED 11:42:55 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 3714 TO ITERATE

53.9% PROCESSED 2000 ITERATIONS 1 ANSWERS  
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 70626 TO 77934

PROJECTED ANSWERS: 1 TO 118

L2 1 SEA SSS SAM L1

=> s l1 sss full

FULL SEARCH INITIATED 11:43:07 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 75735 TO ITERATE

100.0% PROCESSED 75735 ITERATIONS 198 ANSWERS  
SEARCH TIME: 00.00.02

L3 198 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE  
ENTRY

TOTAL  
SESSION

## STN Columbus

FULL ESTIMATED COST

166.94

170.51

FILE 'CAPLUS' ENTERED AT 11:43:17 ON 18 FEB 2006

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FILE LAST UPDATED: 17 Feb 2006 (20060217/ED)

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=> s l3

L4 7 L3

=> d l4 1-7 bib abs fhitr

L4 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

Full Text

AN 2005:980814 CAPLUS

DN 143:431985

TI Privileged structure-based ligands for melanocortin receptors-tetrahydroquinolines, indoles, and aminotetralines

AU Fisher, Matthew J.; Backer, Ryan T.; Husain, Saba; Hsiung, Hansen M.; Mullaney, Jeffrey T.; O'Brian, Thomas P.; Ornstein, Paul L.; Rothhaar, Roger R.; Zgombick, John M.; Briner, Karin

CS Lilly Research Laboratories, A Division of Eli Lilly and Company, Lilly Corporate Center, Indianapolis, IN, 46258, USA

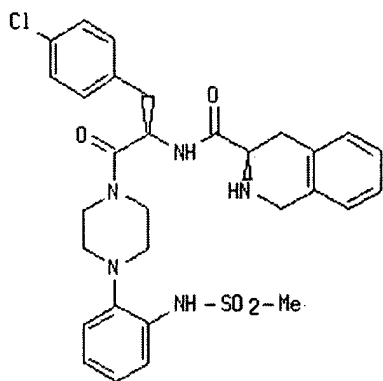
SO Bioorganic Medicinal Chemistry Letters (2005), 15(20), 4459-4462  
CODEN: BMCLE8; ISSN: 0960-894X

PB Elsevier B.V.

DT Journal

LA English

GI



AB Substitution of the aryl sulfonamide moiety contained in MC4 agonist (I) with bicyclic heterocycles and aminotetralines produced compds. with MC4 activity. The heterocycles represent alternative privileged structures to that contained in I. Compds. in which the polar group of the privileged structure was displayed in an endocyclic fashion were not as active as the parent agonist I, while those with an exocyclic polar group afforded activity competitive with I.

IT 444619-63-0P

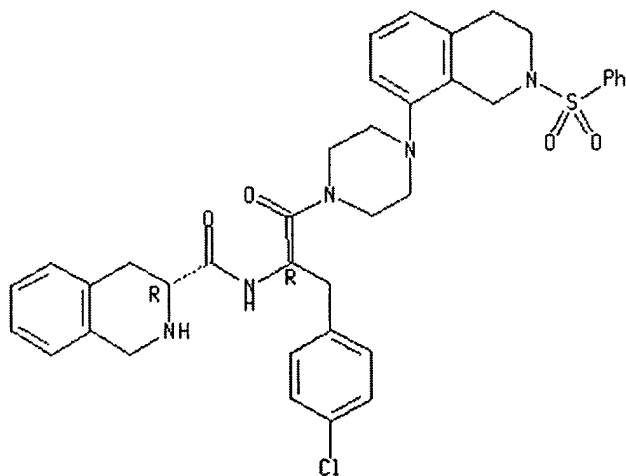
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(privileged structure-based ligands for melanocortin receptors-tetrahydroquinolines, indoles, and aminotetralines)

RN 444619-63-0 CAPLUS

CN 3-Isoquinolinecarboxamide, N-[(1R)-1-[(4-chlorophenyl)methyl]-2-oxo-2-[4-[1,2,3,4-tetrahydro-2-(phenylsulfonyl)-8-isoquinolinyl]-1-piperazinyl]ethyl]-1,2,3,4-tetrahydro-, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



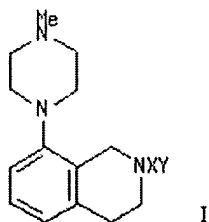
RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
Full Text

## STN Columbus

AN 2004:1080868 CAPLUS  
 DN 142:38280  
 TI Preparation of piperazinyltetrahydroquinolines as serotonin 5-HT1B antagonists.  
 IN Lowe, John Adams, III  
 PA Pfizer Products Inc., USA  
 SO PCT Int. Appl., 23 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

|      | PATENT NO.       | KIND   | DATE     | APPLICATION NO. | DATE     |
|------|------------------|--|----------|-----------------|----------|
| PI   | WO 2004108682    | A2   | 20041216 | WO 2004-IB1855  | 20040601 |
|      | WO 2004108682    | A3   | 20050428 |                 |          |
|      | W:               | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW |          |                 |          |
|      | RW:              | BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |          |                 |          |
|      | US 2004254193    | A1   | 20041216 | US 2004-852726  | 20040524 |
| PRAI | US 2003-477559P  | P  | 20030611 |                 |          |
| OS   | MARPAT 142:38280 |  |          |                 |          |
| GI   |                  |  |          |                 |          |



AB Title compds. [I; X = CH<sub>2</sub>, CH<sub>2</sub>CH<sub>2</sub>, CO, COCH<sub>2</sub>, OCH<sub>2</sub>, OCH<sub>2</sub>CH<sub>2</sub>; Y = alkyl, (substituted) Ph, naphthyl, heteroaryl], were prepd. Thus, 8-bromoisquinoline and PhCH<sub>2</sub>BVr were refluxed together for 6 h in EtOH to give an oil which was stirred with NaBH<sub>3</sub>CN in MeOH for 5 days to give 59% N-benzyl-8-bromotetrahydroisoquinoline. This was heated with N-methylpiperazine, Pd(OAc)<sub>2</sub>, BINAP, and NaOCMe<sub>3</sub> in PhMe at 120° for 16 h to give 84% N-benzyl-8-(4-methylpiperazin-1-yl)tetrahydroisoquinoline. The latter was hydrogenolyzed followed by acylation with 4-chlorophenylacetic acid in the presence of EDAC and N-hydroxybenzotriazole in CH<sub>2</sub>Cl<sub>2</sub> at room temp. to give N-(4-chlorophenylacetyl)-8-(4-methylpiperazin-1-yl)tetrahydroisoquinoline. I showed 5-HT1B receptor affinity with IC<sub>50</sub> <0.60 μM.

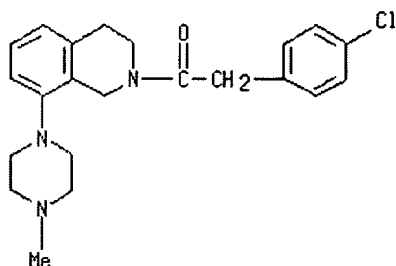
IT 807330-10-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(claimed compd.; prepn. of piperazinyltetrahydroquinolines as serotonin 5-HT1B antagonists)

## STN Columbus

RN 807330-10-5 CAPLUS  
 CN Isoquinoline, 2-[(4-chlorophenyl)acetyl]-1,2,3,4-tetrahydro-8-(4-methyl-1-piperazinyl)- (9CI) (CA INDEX NAME)



L4 ANSWER 3 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

Full Text

AN 2004:756620 CAPLUS

DN 141:277505

TI Preparation of 2,5- and 2,6-substituted tetrahydroisoquinolines as 5-HT6 modulators, in particular selective 5-HT6 antagonists, for treating CNS disorders

IN Putman, David George

PA F. Hoffmann-La Roche A.-G., Switz.

SO PCT Int. Appl., 59 pp.

CODEN: PIXXD2

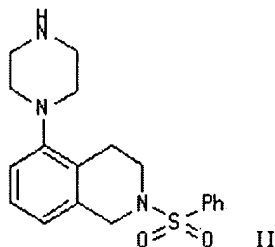
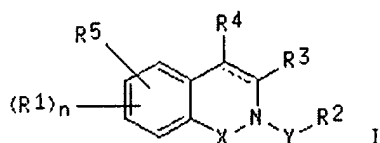
DT Patent

LA English

FAN.CNT 1

|      | PATENT NO.        | KIND | DATE     | APPLICATION NO.  | DATE     |  |
|------|-------------------|------|----------|--|----------|--|
| PI   | WO 2004078176     | A1   | 20040916 | WO 2004-EP1751   | 20040223 |  |
|      | W:                |      |          | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG |          |  |
|      | CA 2517146        | AA   | 20040916 | CA 2004-2517146  | 20040223 |  |
|      | EP 1601358        | A1   | 20051207 | EP 2004-713547   | 20040223 |  |
|      | R:                |      |          | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK   |          |  |
|      | US 2004180874     | A1   | 20040916 | US 2004-791578   | 20040302 |  |
| PRAI | US 2003-451516P   | P    | 20030303 |  |          |  |
|      | WO 2004-EP1751    | W    | 20040223 |  |          |  |
| OS   | MARPAT 141:277505 |      |          |  |          |  |
| GI   |                   |      |          |  |          |  |

APPS



AB Title compds. I [wherein n = 0-3; X = CH<sub>2</sub> and derivs.; C(:O); Y = SO<sub>2</sub> when X = CH<sub>2</sub> and derivs., and Y = (CH<sub>2</sub>)<sub>p</sub> and derivs. when X = C(:O); p = 1-3; each R<sub>1</sub> = independently halo, hetero/halo/alkyl, OH and derivs., NO<sub>2</sub>, CN, S(O)<sub>q</sub>H and derivs., NH<sub>2</sub> and derivs., CONH<sub>2</sub> and derivs., SO<sub>2</sub>NH<sub>2</sub> and derivs., etc.; q = 0-2; R<sub>2</sub> = hetero/aryl, cycloalkyl; R<sub>3</sub>, R<sub>4</sub> = independently H, alkyl; R<sub>5</sub> = 5- to 7-membered nitrogen-contg. heterocycle attached to the 5 or 6 position of the isoquinoline ring; their pharmaceutically acceptable salts or prodrugs] were prepd. as as 5-HT<sub>6</sub> modulators for the treatment of central nervous system (CNS) disorders. Claimed are 5-HT<sub>6</sub> agonists useful for treating CNS diseases. A 5-step synthesis is given for tetrahydroisoquinoline salt II•TFA. In an in vitro radioligand binding study, I were found selective 5-HT<sub>6</sub> antagonists. Selected I showed high affinity for 5-HT<sub>6</sub> receptor with pK<sub>i</sub> values in the range 9.28-10.37 in a radioligand assay. Thus, I are useful in treating mental disorders.

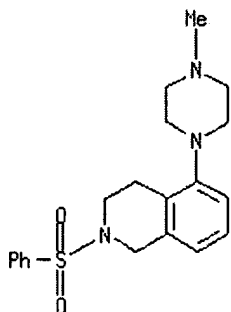
IT 757966-35-1P, 2-Benzenesulfonyl-5-(4-methylpiperazin-1-yl)-1,2,3,4-tetrahydroisoquinoline

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(5-HT<sub>6</sub> antagonist; prepn. of tetrahydroisoquinolines as 5-HT<sub>6</sub> modulators for treating CNS disorders)

RN 757966-35-1 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-5-(4-methyl-1-piperazinyl)-2-(phenylsulfonyl)- (9CI) (CA INDEX NAME)



# STN Columbus

RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

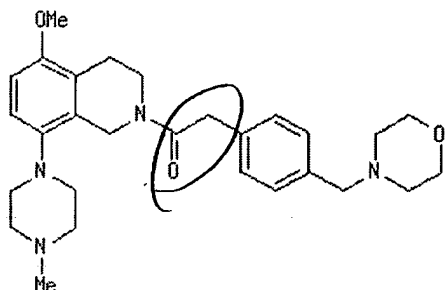
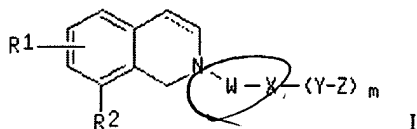
L4 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

## Full Text

AN 2003:356439 CAPLUS  
DN 138:368779  
TI Preparation of isoquinolines as 5-HT antagonists for treatment of  
psychiatric disorders  
IN Angst, Christof; Haeberlein, Markus; Hill, Daniel; Jacobs, Robert; Moore,  
Gary; Pierson, Edward; Shenvi, Ashokkumar Bhikkappa  
PA Astrazeneca AB, Swed.  
SO PCT Int. Appl., 139 pp.  
CODEN: PIXXD2  
DT Patent  
LA English

FAN.CNT 1

|      | PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE     |
|------|---|------|----------|-----------------|----------|
| PI   | WO 2003037887   | A1   | 20030508 | WO 2002-SE1988  | 20021101 |
|      | WO 2003037887   | C1   | 20050317 |                 |          |
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|      | GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, |      |          |                 |          |
|      | LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, |      |          |                 |          |
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|      | UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW                          |      |          |                 |          |
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|      | FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, |      |          |                 |          |
|      | CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG              |      |          |                 |          |
|      | CA 2464342  | AA   | 20030508 | CA 2002-2464342 | 20021101 |
|      | EP 1451172  | A1   | 20040901 | EP 2002-780244  | 20021101 |
|      | R:  |      |          |                 |          |
|      | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, |      |          |                 |          |
|      | IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK          |      |          |                 |          |
|      | BR 2002013778   | A    | 20041109 | BR 2002-13778   | 20021101 |
|      | JP 2005516896   | T2   | 20050609 | JP 2003-540168  | 20021101 |
|      | ZA 2004003240   | A    | 20050407 | ZA 2004-3240    | 20040429 |
|      | NO 2004002154   | A    | 20040729 | NO 2004-2154    | 20040525 |
| PRAI | SE 2001-3644  | A    | 20011101 |                 |          |
|      | WO 2002-SE1988  | W    | 20021101 |                 |          |
| OS   | MARPAT 138:368779   |      |          |                 |          |
| GI   |   |      |          |                 |          |



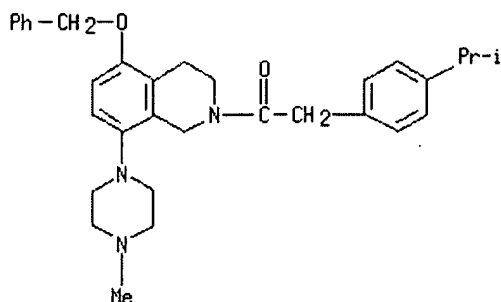
AB Title compds. I [wherein W = CO, CONRa, NRaCO, CO(CH<sub>2</sub>)<sub>n</sub>NRaCO, CSNRa, COCH<sub>2</sub>O, SO<sub>2</sub>NRa, NRaSO<sub>2</sub>, CH<sub>2</sub>NRa, COCH<sub>2</sub>, CH<sub>2</sub>CO, or 5-membered heterocyclyl; X = (un)substituted aryl or heterocyclyl; Y = bond, CH<sub>2</sub>, O, S, SO, CO, SO<sub>2</sub>, NRb, or NRbSO<sub>2</sub>; Z = Rb, CO<sub>2</sub>Ra, CON(Ra)<sub>2</sub>, NHRb, alkyl-N(Ra)<sub>2</sub>, SO<sub>2</sub>Rc, or (un)substituted aryl(alkyl) or heterocyclyl; R1 = halo, alkyl, ORa, SOpRa, N(Ra)<sub>2</sub>, or CN; R2 = aryl or heterocyclyl(carbonyl); Ra = H or (un)substituted alkyl; Rb = H, alkyl(sulfanyl), alkanoyl, aryl(alkyl), or arylalkoxyalkyl; Rc = alkyl, aryl, or heterocyclyl; m = 0 or 1; n = 0-4; p = 0-2;] were prepd. as 5-HT<sub>1B</sub> and 5-HT<sub>1D</sub> antagonists (no data). For example, O-methylation of 5-hydroxyisoquinoline using NaOBu-t and PhMe<sub>3</sub>NC1 in DMF (85%), followed by bromination with bromine in AcOH gave 5-methoxy-8-bromoisquinoline (47%). Substitution with N-methylpiperazine using NaOBu-t, BINAP, and tris(dibenzylideneacetone)dipalladium in PhMe and subsequent redn. with NaCNBH<sub>3</sub> and BF<sub>3</sub>•Et<sub>2</sub>O in MeOH gave 5-methoxy-8-(4-methylpiperazin-1-yl)-1,2,3,4-tetrahydroisoquinoline. Coupling of 4-(bromomethyl)phenylacetic acid with morpholine in the presence of K<sub>2</sub>CO<sub>3</sub> in MeCN provided 4-(morpholinomethyl)phenylacetic acid. Amidation of the tetrahydroisoquinoline with the phenylacetic acid in DMF afforded II. I are useful for the treatment of psychiatric disorders including but not limited to depression, generalized anxiety, eating disorders, dementia, panic disorder, and sleep disorders (no data). The compds. may also be useful in the treatment of gastrointestinal disorders, motor disorders, endocrine disorders, vasospasm, and sexual dysfunction (no data).

IT 521313-56-4P, 1-[5-Benzyloxy-8-(4-methylpiperazin-1-yl)-3,4-dihydro-1H-isoquinolin-2-yl]-2-(4-isopropylphenyl)ethanone  
 RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (5-HT antagonist; prepn. of isoquinolines as 5-HT<sub>1B</sub> and 5-HT<sub>1D</sub> antagonists for treatment of psychiatric disorders)

RN 521313-56-4 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-2-[[4-(1-methylethyl)phenyl]acetyl]-8-(4-methyl-1-piperazinyl)-5-(phenylmethoxy)- (9CI) (CA INDEX NAME)

## STN Columbus



RE.CNT 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

Full Text

AN 2002:575065 CAPLUS

DN 137:140776

TI Preparation of piperidinyl and piperazinyl amino acid derivatives as melanocortin receptor agonists

IN Backer, Ryan Thomas; Briner, Karin; Doecke, Christopher William; Fisher, Matthew Joseph; Kuklish, Steven Lee; Mancuso, Vincent; Martinelli, Michael John; Mullaney, Jeffrey Thomas; Xie, Chaoyu

PA Eli Lilly and Company, USA

SO PCT Int. Appl., 263 pp.

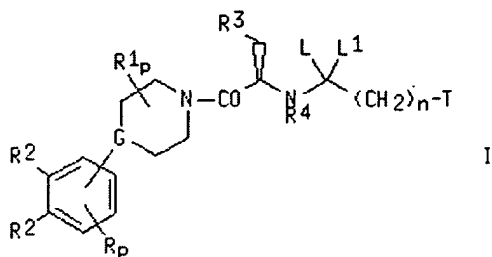
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

|      | PATENT NO.   | KIND | DATE     | APPLICATION NO. | DATE     |
|------|--|------|----------|-----------------|----------|
| PI   | WO 2002059107  | A1   | 20020801 | WO 2002-US516   | 20020123 |
|      | W:   |      |          |                 |          |
|      | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW |      |          |                 |          |
|      | RW:  |      |          |                 |          |
|      | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |      |          |                 |          |
|      | CA 2433025   | AA   | 20020801 | CA 2002-2433025 | 20020123 |
|      | EP 1368339   | A1   | 20031210 | EP 2002-701923  | 20020123 |
|      | R:   |      |          |                 |          |
|      | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR   |      |          |                 |          |
|      | JP 2004521117  | T2   | 20040715 | JP 2002-559409  | 20020123 |
|      | US 2004058936  | A1   | 20040325 | US 2003-466249  | 20030711 |
| PRAI | US 2001-263595P  | P    | 20010123 |                 |          |
|      | WO 2002-US516  | W    | 20020123 |                 |          |
| OS   | CASREACT 137:140776; MARPAT 137:140776   |      |          |                 |          |
| GI   |  |      |          |                 |          |



I

AB The invention relates to melanocortin receptor (MC-R) agonists I [G = CR1 or N; LL1 = H2 or oxo; T = isoquinolinyl or tetrahydro deriv., isoindolinyl, or piperazinyl; n = 0-8; R = H, OH, CN, NO2, halo, alkyl, acyl, etc.; R1 = H, alkyl, alkylcarbamoyl, (D)phenyl, (D)cycloalkyl, or oxo (unless amide is formed); p = 0-4; CR2CR2 is a 5- or 6-membered carbocycle optionally substituted by 1-3 groups R; R3 = (un)substituted aryl or thienyl; R4 = H, alkyl, acyl, cycloalkyl, or alkoxyalkyl], or their pharmaceutically-acceptable salts or stereoisomers, which are useful in the treatment of obesity, diabetes, and male and/or female sexual dysfunction. Comps. I comprise three domains, i.e., a piperidino or piperazinyl fragment, an amino acid, and a radical CLL1(CH2)n-T. Thus, 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid [1-(4-chlorobenzyl)-2-[4-[2-(methylsulfonyl)-1,2,3,4-tetrahydroisoquinolin-8-yl]piperazin-1-yl]-2-oxoethyl]amide (claimed compd.) was prepd. via acylation of the piperazine moiety.

IT 444619-63-0P

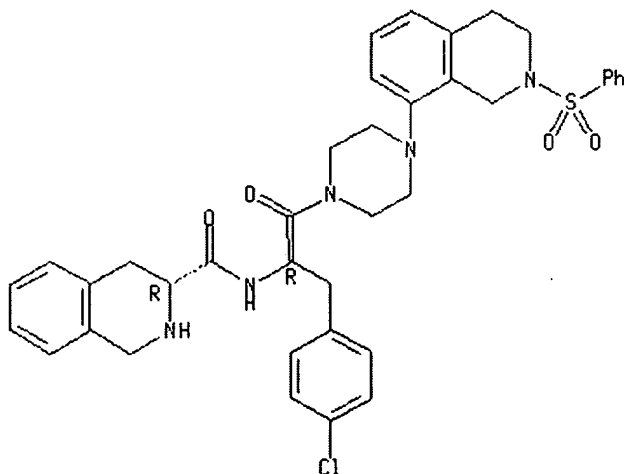
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of piperidiny and piperazinyl amino acid derivs. as melanocortin receptor agonists)

RN 444619-63-0 CAPLUS

CN 3-Isoquinolinecarboxamide, N-[(1R)-1-[(4-chlorophenyl)methyl]-2-oxo-2-[4-[1,2,3,4-tetrahydro-2-(phenylsulfonyl)-8-isoquinolinyl]-1-piperazinyl]ethyl]-1,2,3,4-tetrahydro-, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RE.CNT 6

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

## STN Columbus

L4 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

Full Text

AN 2000:742072 CAPLUS

DN 133:309907

TI Preparation of nitrogen-containing heterocyclic compounds and benzamide compounds as hypolipidemics and antiarteriosclerotics

IN Ohkura, Naoto; Hiraiwa, Yukiko; Matsushima, Tetsuya; Sasaki, Kazue; Yamamoto, Takehiro; Shiotani, Masaharu; Suzuki, Shigeki; Nakatani, Yuuko; Kuroda, Chizuko; Nagasawa, Mieko; Katano, Kiyoaki

PA Meiji Seika Kaisha, Ltd., Japan

SO PCT Int. Appl., 284 pp.

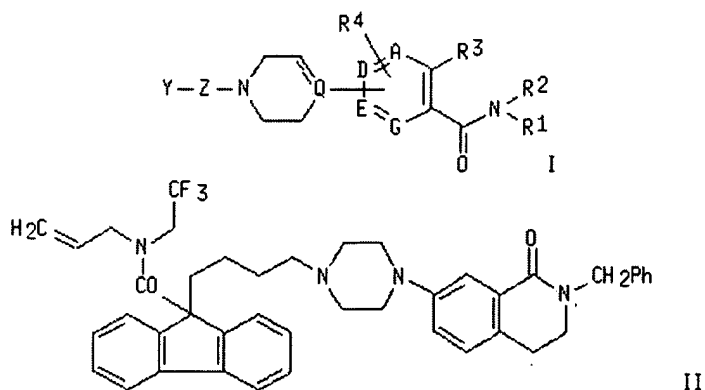
CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 1

|      | PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE     |
|------|---|------|----------|-----------------|----------|
| PI   | WO 2000061556   | A1   | 20001019 | WO 2000-JP2329  | 20000410 |
|      | W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |      |          |                 |          |
|      | RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  |      |          |                 |          |
|      | CA 2369103  | AA   | 20001019 | CA 2000-2369103 | 20000410 |
|      | BR 2000009650   | A    | 20020102 | BR 2000-9650    | 20000410 |
|      | EP 1180514  | A1   | 20020220 | EP 2000-915465  | 20000410 |
|      | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO   |      |          |                 |          |
|      | AU 779550   | B2   | 20050127 | AU 2000-36759   | 20000410 |
|      | US 6777414  | B1   | 20040817 | US 2001-958296  | 20011005 |
|      | US 2004224959   | A1   | 20041111 | US 2004-868006  | 20040616 |
| PRAI | JP 1999-102559  | A    | 19990409 |                 |          |
|      | JP 1999-118490  | A    | 19990426 |                 |          |
|      | JP 1999-119043  | A    | 19990427 |                 |          |
|      | WO 2000-JP2329  | W    | 20000410 |                 |          |
|      | US 2001-958296  | A3   | 20011005 |                 |          |
| OS   | MARPAT 133:309907   |      |          |                 |          |
| GI   |   |      |          |                 |          |



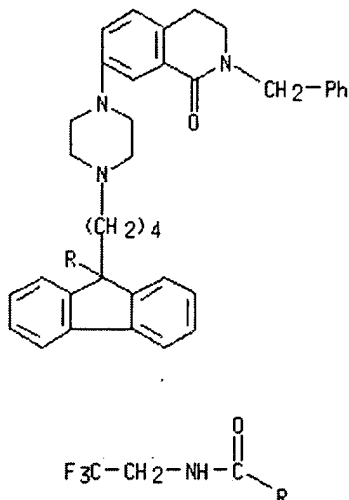
AB The title compds. [I; R1 and R2 represent each (un)substituted C1-6 alkyl or alkoxy, C3-8 cycloalkyl, Ph, C2-6 alkenyl or alkynyl, 5- or 6-membered ring (un)satd. heterocyclyl; R3 and R4 represent each hydrogen, (un)substituted C1-6 alkyl, halo, OH, cyano, C2-5 alkoxy-carbonyl, C1-6 alkoxy, or CO<sub>2</sub>H; or R2 and R3 may be bonded to each other to form (CH<sub>2</sub>)<sub>m</sub>, N:CH, CH:N, or (C1-6 alkyl)-C:N; wherein m is 1 or 2; A, D, E and G represent each C, or one of A, D, E and G represents N and the remainders represent C; Q represents N or C; Y represents a group represented by general formula Q1 (wherein X represents hydrogen, CONR<sub>5</sub>R<sub>6</sub>, etc.; R<sub>8</sub> represents nil or a bond, O, etc.; and R<sub>9</sub> and R<sub>10</sub> represent each hydrogen, alkyl, etc.); and Z represents (CH<sub>2</sub>)<sub>n</sub>, O(CH<sub>2</sub>)<sub>i</sub>, or CONH(CH<sub>2</sub>)<sub>i</sub>; wherein n is 0-6; i is 1-6] are prepd. These compds. have an effect of inhibiting the biosynthesis of triglycerides in the liver and an effect of inhibiting the secretion of apolipoprotein B-contg. lipoproteins from the liver (the latter effect being particularly excellent), without showing the side effect of fat accumulation in the liver, and are useful in treating and preventing hyperlipemia, arteriosclerotic diseases, and pancreatitis. Thus, to a soln. of 2-benzyl-7-[4-[4-[9-(2,2,2-trifluoroethylcarbamoyl)-9H-fluorene-9-yl]butyl]piperazin-1-yl]-3,4-dihydro-2H-isoquinolin-1-one in PhMe were added NaOH, K<sub>2</sub>CO<sub>3</sub>, tetrabutylammonium hydrogen sulfate, and allyl bromide and the resulting mixt. was stirred at 60° overnight to give title compd. (II). II in vitro inhibited the secretion of apolipoprotein B by 89% and the biosynthesis of triglycerides by 89% in HepG2 cells. Tablet and capsule formulations were also described.

IT 301666-52-4P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of nitrogen-contg. heterocyclic compds. and benzamide compds. as hypolipidemics and antiarteriosclerotics and inhibitors of apolipoprotein B-contg. lipoproteins and biosynthesis of triglycerides)

RN 301666-52-4 CAPLUS

CN 9H-Fluorene-9-carboxamide, 9-[4-[4-[1,2,3,4-tetrahydro-1-oxo-2-(phenylmethyl)-7-isoquinolinyl]-1-piperazinyl]butyl]-N-(2,2,2-trifluoroethyl)- (9CI) (CA INDEX NAME)



RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 7 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

Full Text

AN 1998:793126 CAPLUS

DN 130:52434

TI Prepn. of nitrogenous heterocyclic compounds as hyperlipemia remedies

IN Ohkura, Naoto; Tsuruoka, Takashi; Usui, Takayuki; Hiraiwa, Yukiko;  
Matsushima, Tetsuya; Shiotani, Masaharu; Niizato, Tetsutaro; Nakatani,  
Yuuko; Suzuki, Shigeki; Kuroda, Chidsuko; Katano, Kiyooki

PA Meiji Seika Kaisha, Ltd., Japan; et al.

SO PCT Int. Appl., 194 pp.

CODEN: PIXXD2

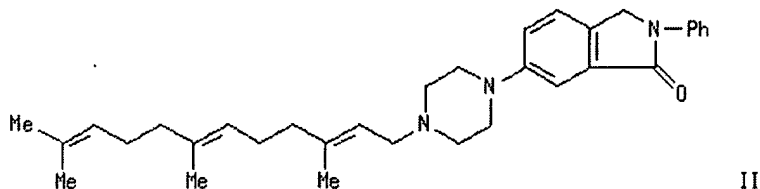
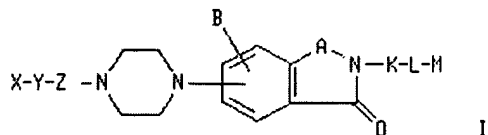
DT Patent

LA Japanese

FAN.CNT 1

|      | PATENT NO.       | KIND   | DATE     | APPLICATION NO. | DATE     |
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| PI   | WO 9854135       | A1   | 19981203 | WO 1998-JP2411  | 19980601 |
|      | W:               | AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |          |                 |          |
|      | RW:              | GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG   |          |                 |          |
|      | CA 2291630       | AA   | 19981203 | CA 1998-2291630 | 19980601 |
|      | AU 9875482       | A1   | 19981230 | AU 1998-75482   | 19980601 |
|      | EP 999208        | A1   | 20000510 | EP 1998-923066  | 19980601 |
|      | R:               | DE, ES, FR, GB, IT   |          |                 |          |
|      | US 6417362       | B1   | 20020709 | US 1999-424708  | 19991130 |
|      | US 2002156276    | A1   | 20021024 | US 2002-127491  | 20020423 |
|      | US 6583144       | B2   | 20030624 |                 |          |
| PRAI | JP 1997-141410   | A  | 19970530 |                 |          |
|      | WO 1998-JP2411   | W  | 19980601 |                 |          |
|      | US 1999-424708   | A3   | 19991130 |                 |          |
| OS   | MARPAT 130:52434 |  |          |                 |          |
| GI   |                  |  |          |                 |          |

# STN Columbus



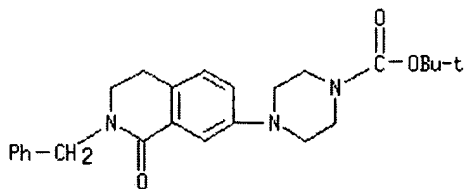
AB The title compds. [I; A = CR<sub>1</sub>R<sub>2</sub>(CH<sub>2</sub>)<sub>i</sub>; (wherein R<sub>1</sub> and R<sub>2</sub> each represents hydrogen or alkyl, i = 0-1), CH:CH, OCH<sub>2</sub>, or S(O)<sub>j</sub>CH<sub>2</sub> (wherein j = 0-2); B = hydrogen or halogen; X = CR<sub>3</sub>R<sub>4</sub>R<sub>5</sub>, NR<sub>6</sub>R<sub>7</sub>, (CH<sub>2</sub>CH:C(CH<sub>3</sub>)CH<sub>2</sub>)<sub>p</sub>CH<sub>2</sub>CH:C(CH<sub>3</sub>)<sub>2</sub>, alkyl, cycloalkyl, Ph, cinnamyl, or heteroaryl; Y = (CH<sub>2</sub>)<sub>q</sub>, CH:CH, NR<sub>8</sub>, oxygen, or a bond; Z = carbonyl or a bond; K = alkylene or a bond; L = CH:CH or a bond; and M = hydrogen, alkyl, cycloalkyl, Ph, heterocycle, biphenyl, or diphenylmethyl; p = 0-2; q = 1-6; R<sub>3</sub>-R<sub>5</sub> = hydrogen, phenyl; R<sub>6</sub>-R<sub>7</sub> = hydrogen, Ph, benzyl; R<sub>8</sub> = hydrogen, C1-6 alkyl] are prepd. I inhibit the biosynthesis of triglycerides in the liver and also inhibit the secretion of lipoproteins contg. apolipoprotein B from the liver. I are hence useful for the prevention/treatment of hyperlipemia (esp. hyper-VLDL-emia) and diseases caused thereby, such as arteriosclerotic diseases, e.g., myocardial infarct, and pancreatitis. Thus, title compd. (II) was prepd. by multi-step reactions and showed 56% and 90% inhibitory activity for apolipoprotein B and triglycerides. A formulation contg. I was also presented.

IT 217492-19-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of nitrogenous heterocyclic compds. as hyperlipemia remedies)

RN 217492-19-8 CAPLUS

CN 1-Piperazinecarboxylic acid, 4-[1,2,3,4-tetrahydro-1-oxo-2-(phenylmethyl)-7-isoquinolinyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



RE.CNT 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file caold

COST IN U.S. DOLLARS

| SINCE FILE | TOTAL   |
|------------|---------|
| ENTRY      | SESSION |
| 38.07      | 208.58  |

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

| SINCE FILE | TOTAL   |
|------------|---------|
| ENTRY      | SESSION |

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FILE LAST UPDATED: 01 May 1997 (19970501/UP)

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| COST IN U.S. DOLLARS                       | SINCE FILE | TOTAL   |
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\* The CA roles and document type information have been removed from \*  
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\* available and contains the CA role and document type information. \*  
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Structure search iteration limits have been increased. See HELP SLIMITS for details.

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<http://www.cas.org/ONLINE/UG/regprops.html>

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FILE 'REGISTRY' ENTERED AT 11:42:45 ON 18 FEB 2006

L1 STRUCTURE UPLOADED

L2 1 S L1

L3 198 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 11:43:17 ON 18 FEB 2006

L4 7 S L3

FILE 'CAOLD' ENTERED AT 11:46:07 ON 18 FEB 2006

L5 0 S L3

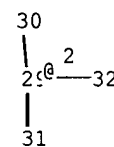
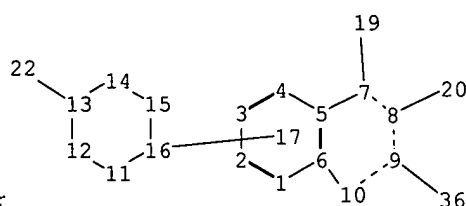
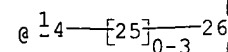
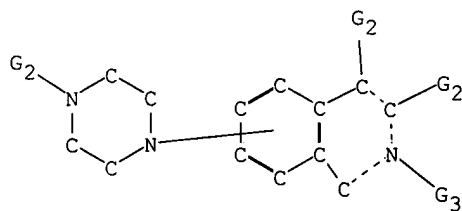
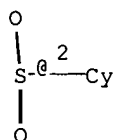
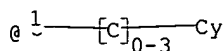
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chain nodes :

19 20 22 24 25 26 29 30 31 32 36

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16

chain bonds :

7-19 8-20 9-36 13-22 24-25 25-26 29-30 29-31 29-32

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-10 7-8 8-9 9-10 11-12 11-16  
12-13 13-14 14-15 15-16

exact/norm bonds :

5-7 6-10 7-8 7-19 8-9 8-20 9-10 9-36 11-12 11-16 12-13 13-14  
13-22 14-15 15-16 25-26 29-30 29-31 29-32

exact bonds :

24-25

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6

isolated ring systems :

containing 1 : 11 :

G2:C,H

G3:[\*1],[\*2]

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom  
10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom  
19:CLASS 20:CLASS 22:CLASS 24:CLASS 25:CLASS 26:Atom 29:CLASS  
30:CLASS

31:CLASS 32:Atom 36:CLASS